SEARCH REQUEST FORM

Scientific and Technical Information Center

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Requester's Full Name: Notel		Examiner #: 78 462 Date:	<u> </u>
	Number 30	Serial Number: <u>09/589, 777</u> Results Format Preferred (circle): PAPER DISK I	 7-M A II
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nclude the elected species or structures,	keywords, synonyms, a	cronyms, and registry numbers, and combine with the conc	ept or
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nventors (please provide full names):		 	
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Earliest Priority Filing Date: 4-	22-98		
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Davis, N. 09/589777

09/589777

FILE 'REGISTRY' ENTERED AT 10:57:57 ON 06 APR 2001 17 S SYIVLCIE/SQSP

FILE 'CAPLUS' ENTERED AT 10:58:36 ON 06 APR 2001 19 S L1

ANSWER 1 OF 19 CAPLUS COPYRIGHT 2001 ACS L2

ACCESSION NUMBER:

2001:208854 CAPLUS

DOCUMENT NUMBER:

134:217893

TITLE:

L1

L2

Functional annotation of a full-length mouse

cDNA collection

AUTHOR (S):

Kawai, J.; Shingawa, A.; Shibata, K.; Yoshino, M.; Itoh, M.; Ishii, Y.; Arakawa, T.; Hara, A.; Funkunishi, Y.; Konno, H.; Adachi, J.; Fukuda, S.; Aizawa, K.; Izawa, M.; Nishi, K.; Kiyosawa, H.; Kondo, S.; Yamanaka, I.; Saito, T.; Okazaki, Y.; Gojobori, T.; Bono, H.; Kasukawa, T.; Saito, R.; Kadota, K.; Matsuda, H.; Ashburner, M.; Batalov, S.; Casavant, T.; Fleischmann, W.; Gaasterland, T.; Gissi, C.; King, B.; Kochiwa, H.; Kuehl, P.; Lewis, S.; Matsuo, Y.; Nikaido, I.; Pesole, G.; Quackenbush, J.; Schriml, L. M.; Staubli, F.; Suzuki, R.; Tomita, M.; Wagner, L.; Washio, T.; Sakai, K.; Okido, T.; Furuno, M.; Aono, H.; Baldarelli, R.; Barsh, G.; Blake, J.; Boffelli, D.; Bojunga, N.; Carninci, P.; de Bonaldo, M. F.; Brownstein, M. J.; Bult, C.; Fletcher, C.; Fujita, M.; Gariboldi, M.; Gustincich, S.; Hill, D.; Hofmann, M.; et al. Lab. Genome Exploration Res. Group, RIKEN

CORPORATE SOURCE:

Genomic Sciences Center (GSC), Yokohama Inst.,

308-4994

Yokohama, kanagawa, 230-0045, Japan

SOURCE:

Nature (London) (2001), 409(6821), 685-690

CODEN: NATUAS; ISSN: 0028-0836

PUBLISHER:

Nature Publishing Group

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB The RIKEN Mouse Gene Encyclopaedia Project, a systematic approach to detg. the full coding potential of the mouse genome, involves collection and sequencing of full-length cDNAs and phys. mapping of the corresponding genes to the mouse genome. An international functional annotation meeting (FANTOM) was organized to annotate the first 21,076 cDNAs to be analyzed in this project. This report describes the first RIKEN clone collection, which is one of the largest described for any organism. Anal. of these cDNAs extends known gene families and identifies new ones. The sequences are deposited into GenBank with Accession nos. AK002213-AK021412 and AK027261-AK027262. Information about these clones is available at RIKEN (http://www.gsc.riken.go.jp/e/FANTOM/viewer/) and Mouse Genome Searcher Shears

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Informatics (http://www.informatics.jax.org and mirror sites). [This abstr. record is one of 7 records for this document necessitated by the large no. of index entries required to fully index the document and publication system constraints.]. 326629-20-3

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; functional annotation of a full-length mouse cDNA collection)

ANSWER 2 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

2001:154589 CAPLUS

TITLE:

IT

Functional annotation of a full-length mouse

cDNA collection

AUTHOR (S):

Kawai, J.; Shingaawa, A.; Shibata, K.; Yoshino, M.; Itoh, M.; Ishii, Y.; Aarakawa, T.; Hara, A.; Fukunishi, Y.; Konno, H.; Adcahi, J.; Fukuda, S.; Aizawa, K.; Izawa, M.; Nishi, K.; Kiyosawa, H.; Kondo, S.; Yamanaka, I.; Saito, T.; Okazaki, Y.; Gojobori, T.; Bono, H.; Kasukawa, T.; Saito, R.; Kadota, K.; Matsuda, H.; Ashburner, M.; Batalov, S.; Csavant, T.; Fleischmann, W.; Gaasterland, T.; Gissi, C.; King, B.; Kochiwa, H.; Kuehl, P.; Lewis, S.; Matsuo, Y.; Nikaido, I.; Pesole, G.; Quackenbush, J.; Schriml, L. M.; Staubli, F.; Suzuki, R.; Wagner, L.; Wagner, L.; Washio, T.; Sakai, K.; Okido, T.; Furuno, M.; Aono, H.; Baldarelli, R.; Barsh, G.; Blake, J.; Boffelli, D.; Bojunga, N.; Carninci, P.; de Bonaldo, M. F.; Brownstein, M. J.; Bult, C. fletcher, C.; Fujita, M.; Gariboldi, M.; Gustincichh, S.; Hill, D.; Hofmann, M.; Hume, D. A.; Kamiya, M.; Lee, N. H.; Lyons, P.; Marchionni, L.; Mashima, J.; Mazzarelli, J.; Mombaerts, P.; Nordone, P.; Ring, B.; Ringwald, M.; Mombaerts, P.; Rodriquez, I.; Sakamoto, N.; Sasaki, H.; Sato, K.; Schonbach, C.; Seya, T.; Shibata, Y.; Storch, K.-F.; Suzuki, H.; Toyo-oka, K.; Wang, K. H.; Weitz, C.; Whittaker, C.; Wilming, L.; Wynshaw-Boris, A.; Yoshida, K.; Hasegawa, Y.; Kawaji, H.; Kohtsuki, S. Lab. Genome Explortaion Res. Group, RIKEN Genomic Sciences Center (GSC), Yokohama Inst.,

CORPORATE SOURCE:

Yokohama, Kanagawa, 230-0045, Japan

Nature (London) (2001), 409(6821), 685-690

CODEN: NATUAS; ISSN: 0028-0836

Nature Publishing Group

Journal

DOCUMENT TYPE:

English

LANGUAGE:

PUBLISHER:

SOURCE:

AB The RIKEN Mouse Gene Encyclopaedia Project, a systematic approach to detg. the full coding potential of the mouse genome, involves collection and sequencing of full-length cDNAs and phys. mapping of the corresponding genes to the mouse genome. An international functional annotation meeting (FANTOM) was organized to annotate the first 21,076 cDNAs to be analyzed in this project. This report describes the first RIKEN clone collection, which is one of the largest described for any organism. Anal. of these cDNAs extends known gene families and identifies new ones. The sequences are deposited into GenBank with Accession nos. AK002213-AK021412 and AK027261-AK027262. Information about these clones is available at RIKEN (http://www.gsc.riken.go.jp/e/FANTOM/viewer/) and Mouse Genome Informatics (http://www.informatics.jax.org and mirror sites). [This abstr. record is one of 7 records for this document necessitated by the large no. of index entries required to fully index the document and publication system constraints.].

IT 326629-20-3

RL: BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study)

(amino acid sequence; functional annotation of a full-length mouse cDNA collection)

L2 ANSWER 3 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2001:137393 CAPLUS

DOCUMENT NUMBER: 134:188970

TITLE: Adenoviral vectors including DNA sequences

encoding angiogenic inhibitors

INVENTOR(S): Hallenbeck, Paul L.; Chen, Cheauyun Theresa

PATENT ASSIGNEE(S): Novartis A.-G., Switz.; Novartis-Erfindungen

Verwaltungsgesellschaft m.b.H.

SOURCE: PCT Int. Appl., 59 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PAT	rent 1	NO.		KI.	ND	DATE			A	PPLI	CATI	ON NO	ο.	DATE		
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WO	2001	0128	30	Α	1	2001	0222		W	20	00-E	P786	5	2000	0811	
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		CN,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KP,	KR,	KZ,	LC,	LK,
		LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,
		PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,
		UA,	ŪĠ,	UZ,	VN,	YU,	ZA,	ZW,	AM,	AZ,	BY,	KG,	KZ,	MD,	RU,	TJ,
		TM														
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	AT,	BE,	CH,
		CY,	DE,	DK.	ES,	FI,	FR.	GB,	GR,	IE,	IT,	LU,	MC.	NL,	PT,	SE,
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BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.: US 1999-373938 19990813

AB An adenoviral vector which includes at least one DNA sequence encoding an angiogenic inhibitor, such as endostatin. Such vectors may be employed in treating diseases or disorders assocd. with angiogenesis, such as cancer, vascular diseases of the eye, including diabetic retinopathy, psoriasis, arthritis, cardiovascular disease, cerebral edema and Kasabach-Merritt syndrome. Recombinant adenoviral vectors encoding murine (or human endostatin) are constructed for making transgenic mice. Expression and secretion of murine endostatin and the functions of the secreted endostatin are studied from various disease models, including colon liver metastasis model, B16F10 lung metastasis model, B16F10 s.c. model, B16F10 melanoma lung metastasis model, and B16F10 melanoma s.c. model.

IT 326948-44-1

RL: PRP (Properties)

(unclaimed protein sequence; adenoviral vectors including DNA sequences encoding angiogenic inhibitors)

REFERENCE COUNT:

REFERENCE(S):

- (1) Blezinger, P; NATURE BIOTECHNOLOGY 1999, V17(4), P343 CAPLUS
- (2) Crystal, R; NATURE BIOTECHNOLOGY 1999, V17, P336 CAPLUS
- (3) Feldman, A; WO 0068379 A 2000 CAPLUS
- (4) Genetix Pharmaceuticals Inc; WO 9926480 A 1999 CAPLUS

Shears

308-4994

(5) LI, H; WO 9849321 A 1998 CAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 4 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:814324 CAPLUS

DOCUMENT NUMBER: 134:505

TITLE: Antiangiogenic endostatin peptides, endostatin

variants and methods of use

INVENTOR(S):
Vuori, Kristiina

PATENT ASSIGNEE(S): The Burnham Institute, USA SOURCE: PCT Int. Appl., 146 pp.

CODEN: PIXXD2

Searcher

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2000067771 A1 20001116 WO 2000-US12063 20000502

W: AE, AG, AL, AM, AT, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH,

CN, CR, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EE, EE, ES, FI,

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FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG,
             KP, KR, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK,
             MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK,
             SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                                                            19990506
                                           US 1999-132907
PRIORITY APPLN. INFO.:
                                           US 1999-353333
                                                             19990714
     The invention provides an endostatin peptide having at least 4-7
AB
     endostatin amino acid residues contg. substantially the amino acid
     sequence of RLQD, RAD, DGK/R, or a functional equiv. thereof. The
     invention also provides an endostatin variant contg. the amino acid
     sequence RGD, or a functional fragment thereof. Methods of
     inhibiting angiogenesis are also provided.
     307924-80-7
IT
     RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological
     study); OCCU (Occurrence)
        (antiangiogenic endostatin peptides, endostatin variants and
        methods of use)
     224308-23-0
IT
     RL: PRP (Properties)
        (unclaimed protein sequence; antiangiogenic endostatin peptides,
        endostatin variants and methods of use)
REFERENCE COUNT:
                         (1) Brooks; US 5753230 A 1998 CAPLUS
REFERENCE(S):
                         (2) Koivunen, E; Journal of Biological Chemistry
                             1993, V268(27), P20205 CAPLUS
                         (3) La Jolla Cancer Research Foundation; WO
                             9514714 A1 1995 CAPLUS
                         (4) Nutt; US 5061693 A 1991 CAPLUS
                         (5) Oh, S; Proc Natl Acad Sci USA 1994, V91,
                             P4229 CAPLUS
                         ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 5 OF 19 CAPLUS COPYRIGHT 2001 ACS
L2
                         2000:772677 CAPLUS
ACCESSION NUMBER:
                         133:349140
DOCUMENT NUMBER:
                         Compositions and methods for cancer treatment by
TITLE:
                         selectively inhibiting VEGF
                         Thorpe, Philip E.; Brekken, Rolf A.
INVENTOR (S):
                         Board of Regents, the University of Texas
PATENT ASSIGNEE(S):
                         System, USA
                         PCT Int. Appl., 297 pp.
SOURCE:
                         CODEN: PIXXD2
                         Patent
DOCUMENT TYPE:
                         English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
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PATENT INFORMATION:

	IND DATE APPLICATION NO. DATE
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WO 2000064946	
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	, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,
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	, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT,
	, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,
	, 3E, 3G, 3I, 3K, 3E, 10, 1M, 1K, 11, 12, 0A, 0G, 2A, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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	, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF,
	, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:	US 1999-131432 19990428
	bodies that specifically inhibit VEGF binding to
	of the two VEGF receptors. The antibodies
	t angiogenesis and induce tumor regression, and
	safety due to their specificity. The present
	vides new antibody-based compns., methods and
-	for treating cancer and other angiogenic
	geous immunoconjugate and prodrug compns. and
methods using the i	new VEGF-specific antibodies are also provided.
IT 304489-40-5DP, imm	unoconjugates
RL: BPN (Biosynthet	tic preparation); PRP (Properties); THU
(Therapeutic use);	BIOL (Biological study); PREP (Preparation); USES
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(amino acid com	uence; immunoconjugates of anti-VEGF antibody for
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diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE:	herapy of cancer and angiogenic disease) PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity
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diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy;
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S):	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp.
diagnosis and the L2 ANSWER 6 OF 19 CANACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent English
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent English
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent English
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent English 1
diagnosis and the L2 ANSWER 6 OF 19 CAN ACCESSION NUMBER: DOCUMENT NUMBER: TITLE: INVENTOR(S): PATENT ASSIGNEE(S): SOURCE: DOCUMENT TYPE: LANGUAGE: FAMILY ACC. NUM. COUNT:	PLUS COPYRIGHT 2001 ACS 2000:756746 CAPLUS 133:329588 Endostatin-derived peptides exhibiting antiangiogenic activity Chillemi, Francesco; Francescato, Pierangelo; Ziche, Marina Universita' Degli Studi Di Milano, Italy; Universita' Degli Studi Di Firenze PCT Int. Appl., 28 pp. CODEN: PIXXD2 Patent English 1

WO 2000063249 A1 20001026 WO 2000-EP3236 20000411 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG PRIORITY APPLN. INFO.: IT 1999-MI777 19990415 Peptides with a sequence corresponding or homologous to that of endostatin, having inhibiting activity on angiogenesis, are useful in the treatment of angiogenesis-dependent tumors. 303113-25-9P ΙT RL: BAC (Biological activity or effector, except adverse); PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (endostatin-derived peptides with antiangiogenic activity) IT 303042-57-1DP, resin-bound RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation) (prepn. and reaction; endostatin-derived peptides with antiangiogenic activity) REFERENCE COUNT: REFERENCE(S): (1) Beth Israel Deaconess Medical Center; WO 9929855 A 1999 CAPLUS (2) The Children'S Medical Center Corporation; WO 9715666 A 1997 CAPLUS (3) The Children'S Medical Center Corporation; EP 0857210 A 1998 CAPLUS (4) The Children'S Medical Center Corporation & Yissum Research; WO 9948924 A 1999 CAPLUS ANSWER 7 OF 19 CAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 2000:707320 CAPLUS DOCUMENT NUMBER: 133:292674 Soluble recombinant endostatin TITLE: INVENTOR(S): Boice, Judith A.; Leiting, Barbara; O'Connell, John F.; Pompliano, David L. PATENT ASSIGNEE(S): Merck & Co., Inc., USA SOURCE: PCT Int. Appl., 58 pp. CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: English FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE

Searcher

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308-4994

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WO 2000058498 A1 20001005
                                          WO 2000-US8435 20000329
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN,
             CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS,
             LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO,
             RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
             UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
         RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF,
             BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                           US 1999-126806 19990330
                                           US 1999-128489 19990409
AB
     A method of making sol. recombinant endostatin in Escherichia coli
     is disclosed. Prepns. contg. the sol. recombinant endostatin are
     provided for the inhibition of angiogenesis or the treatment of
     cancer. Assays using the sol. recombinant endostatin are also
     provided. Thus, the structure of sol. endostatin prepd. by the
     method of the invention was detd. by NMR. The data indicated that
     endostatin exists in a monomeric form in soln. The NMR structure of
     zinc-contg. endostatin shows that the N.delta. imidazole nitrogen
     atoms of residues His-1, His-3, and His-11 are chelating the Zn2+.
ΙT
     259789-72-5P, Endostatin (mouse)
     RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation);
     PUR (Purification or recovery); BIOL (Biological study); PREP
     (Preparation)
        (sol. recombinant endostatin)
REFERENCE COUNT:
REFERENCE(S):
                         (1) Cohen, J; Science 1999, V283, P1250 CAPLUS
                         (2) Dhanabal; Cancer Research 1999, V59, P189
                             CAPLUS
                         (3) G D Searle & Co; WO 9942486 A1 1999 CAPLUS
                         (4) Kohno; Methods in Enzymology 1990, V185,
                             P187 CAPLUS
                         (5) Lowe; Solubilisation, refolding and
                             purification of eukaryotic proteins
                             expressed in E coli in Protein purification
                             1987, P429 CAPLUS
                         ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 8 OF 19 CAPLUS COPYRIGHT 2001 ACS
                        2000:434233 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                        133:79332
TITLE:
                        Carrier-DNA complexes containing DNA encoding
                         anti-angiogenic peptides and their use in gene
                        therapy
INVENTOR (S):
                        Mixson, A. James
```

U.S., 30 pp., Cont.-in-part of U.S. 5,815,216.

: Shears 308-4994

PATENT ASSIGNEE(S):

SOURCE:

USA

Searcher

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

KIND DATE	APPLICATION NO.	DATE
A 20000627	US 1997-985526	19971205
A2 19980121	EP 1997-112154	19970716
A3 19980204		
CH, DE, DK, ES, FR,	GB, GR, IT, LI, LU,	, NL, SE, MC,
SI, LT, LV, FI, RO		
A1 19990609	EP 1998-100135	19980107
CH, DE, DK, ES, FR,	GB, GR, IT, LI, LU,	NL, SE, MC,
SI, LT, LV, FI, RO		
A2 19990713	JP 1998-201996	19980716
.:	US 1996-680845	19960716
	EP 1997-112154	19970716
	US 1997-985526	19971205
	A 20000627 A2 19980121 A3 19980204 CH, DE, DK, ES, FR, SI, LT, LV, FI, RO A1 19990609 CH, DE, DK, ES, FR, SI, LT, LV, FI, RO A2 19990713	A 20000627 US 1997-985526 A2 19980121 EP 1997-112154 A3 19980204 CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, SI, LT, LV, FI, RO A1 19990609 EP 1998-100135 CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, SI, LT, LV, FI, RO A2 19990713 JP 1998-201996 .: US 1996-680845 EP 1997-112154

AB Carrier complexes comprising DNA encoding an anti-angiogenic gene or peptide and optionally a further DNA encoding a tumor suppressor protein are described. When administered to a subject bearing a tumor, the complexes can inhibit growth of the tumor.

IT 226938-38-1, Endostatin (human fragment)

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(amino acid sequence; carrier-DNA complexes contg. DNA encoding anti-angiogenic peptides and their use in gene therapy)

REFERENCE COUNT:

40

REFERENCE(S):

(1) Anon; EP 0443404 A1 1991 CAPLUS (2) Anon; WO 9202240 1992 CAPLUS (3) Anon; WO 9316716 1993 CAPLUS (4) Anon; WO 9316718 1993 CAPLUS (5) Anon; WO 9529242 1995 CAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 9 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

2000:144909 CAPLUS

DOCUMENT NUMBER:

132:190503

TITLE:

Expression and export of angiostatin and

endostatin as immunofusins

INVENTOR (S):

Lo, Kin-Ming; Li, Yue; Gillies, Stephen D.

PATENT ASSIGNEE(S): Lexingen Pharmaceuticals Corp., USA

SOURCE:

PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE: EN FAMILY ACC. NUM. COUNT: 1

English

PATENT INFORMATION:

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KIND DATE
     PATENT NO.
                                         APPLICATION NO. DATE
     ______
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                          -----
                                          -----
     WO 2000011033
                           20000302
                                          WO 1999-US19329 19990825
                      A2
     WO 2000011033
                     A3
                           20000622
        W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
            CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
            ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD,
            SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA,
            ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE,
            DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
            CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
     AU 9955836
                      A1 20000314
                                         AU 1999-55836
                                                          19990825
PRIORITY APPLN. INFO.:
                                         US 1998-97883
                                                          19980825
                                         WO 1999-US19329 19990825
    Disclosed are nucleotide sequences, for example, DNA or RNA
AB
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sequences, which encode an Ig Fc-angiogenesis inhibitor fusion protein. The angiogenesis inhibitors can be angiostatin, endostatin, a plasminogen fragment having angiostatin activity, or a collagen XVIII fragment having endostatin activity. The nucleotide sequences can be inserted into a suitable expression vector and expressed in mammalian cells. Also disclosed is a family of Ig Fc-angiogenesis inhibitor fusion proteins that can be produced by expression of such nucleotide sequences. Also disclosed are methods using such nucleotide sequences and fusion proteins for treating conditions mediated by angiogenesis. When C57/BL6 mice with implanted Lewis lung tumors are injected with 720 .mu.g human Fc-human angiostatin fusion protein per mouse, the protein had a circulating half-life of about 32 h, and Western anal. shows that >90% of the fusion protein remains as an intact mol. in circulation. 259789-72-5DP, Endostatin (mouse), fusion products ΙT

RL: BPN (Biosynthetic preparation); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(amino acid sequence; expression and export of angiostatin and endostatin as immunofusins)

L2 ANSWER 10 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:62891 CAPLUS

DOCUMENT NUMBER: 132:103744

TITLE: Cloning of cDNA for human endostatin and use for

inhibition of angiogenesis

INVENTOR(S): Xu, Genxing; Ren, Mindong; Xu, Lin

PATENT ASSIGNEE(S): Peop. Rep. China

SOURCE: Faming Zhuanli Shenqing Gongkai Shuomingshu, 6

pp.

CODEN: CNXXEV

DOCUMENT TYPE:

Patent

LANGUAGE:

Chinese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
CN 1177005	A	19980325	CN 1997-107112	19970910
CN 1060521	R	20010110		

Described is a method of cloning the cDNA for human liver endostatin AB by PCR using a pair of primers derived from the cDNA encoding human collagen type XVIII (1503-2055 cDNA fragment). Endostatin is useful for the treatment of tumors by inhibiting angiogenesis.

255811-03-1 ΙT

RL: PRP (Properties)

(unclaimed sequence; cloning of cDNA for human endostatin and use for inhibition of angiogenesis)

ANSWER 11 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1999:388287 CAPLUS

DOCUMENT NUMBER:

131:41277

TITLE:

Mutants of endostatin, "em 1" having

anti-angiogenic activity and methods of use

thereof

INVENTOR(S):

Sukhatme, Vikas P.

PATENT ASSIGNEE(S):

Beth Israel Deaconess Medical Center, USA

SOURCE:

PCT Int. Appl., 105 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PA'	rent :	NO.		KI	ND	DATE			A	PPLI	CATI	ON NO	ο.	DATE		
									-		- -					
WO	9929	855		A	1	1999	0617		W	0 19	98-U	S260	57	1998	1208	
	W:	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,
		DE,	DK,	ΕĒ,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	ΗU,	ID,	IL,	IN,
		IS,	JP,	KΕ,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,
		MG,	MK,	MN,	MW,	MX,	NO,	NZ,	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,
		SK,	SL,	TJ,	TM,	TR,	TT,	UA,	ŪĠ,	US,	US,	US,	UΖ,	VN,	YU,	ZW,
		AM,	ΑZ,	BY,	KG,	KZ,	MD,	RU,	ТJ,	TM						
	RW:	GH,	GM,	ΚE,	LS,	MW,	SD,	SZ,	υG,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,
		ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,
		CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG			
	9917					1999										
EP	1037															
	R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,
					Se	arch	er	:		She	ars	3	08-4	994		

PT, IE, FI

PRIORITY APPLN. INFO.:

US 1997-67888 19971208 US 1998-82663 19980422 US 1998-108536 19981116 WO 1998-US26057 19981208

AB Described herein are novel mutants of endostatin, one of which, designated "EM 1", has anti-angiogenic activity similar or superior to that of wild type endostatin. The invention relates to the discovery of an isolated anti-angiogenic peptide, wherein the C-terminal end of the peptide comprises the amino acid sequence SYIVLCIE, which has anti-angiogenic properties. Designated "EM 1", this protein comprises a mutated endostatin protein, where the mutation comprises a deletion of nine consecutive amino acids from the C-terminus of the mutated endostatin protein (e.g., NSFMTSFSK). EM 1 terminates in the amino acid sequence SYIVLCIE. The invention also comprises isolated polynucleotides encoding EM 1, operably linked to expression sequence, and host cells transformed with such a construct. Antibodies to EM 1 are also disclosed. The invention also relates to processes for producing EM 1, fusion proteins contg. EM 1, and compns. comprising EM 1 or fusion products thereof. invention also discloses methods of producing polypeptides encoding EM 1.

IT 224308-23-0

RL: BAC (Biological activity or effector, except adverse); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(mutants of endostatin, "em 1" having anti-angiogenic activity and methods of use thereof)

REFERENCE COUNT:

8

REFERENCE(S):

- (1) Boehm, T; Biochemical and Biophysical Research Communications 1998, V252, P190 CAPLUS
- (2) Dhanabal, M; Cancer Research 1999, V59, P189 CAPLUS
- (3) Ding, Y; Proc Natl Acad Sci USA 1998, V95, P10443 CAPLUS
- (5) Hohenester, E; The EMBO Journal 1998, V17(6), P1656 CAPLUS
- (7) O'Reilly, M; Cell 1997, V88(2), P277 CAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 12 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1999:375339 CAPLUS

DOCUMENT NUMBER:

131:28626

TITLE:

Delivery of anti-angiogenic genes to a tumor in

vivo and their use in gene therapy

INVENTOR(S):

Mixson, Archibald James

PATENT ASSIGNEE(S):

USA

SOURCE:

Eur. Pat. Appl., 46 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

EP 921193 A1 19990609 EP 1998-100135 19980107

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC,

PT, IE, SI, LT, LV, FI, RO

US 6080728 A 20000627 PRIORITY APPLN. INFO.:

US 1997-985526 19971205 US 1997-985526 19971205

US 1996-680845 19960716

EP 1997-112154 19970716

AB The invention relates to the delivery of anti-angiogenic genes or DNA encoding anti-angiogenic peptides to a tumor in vivo, preferably by injection, and expression of the DNA in order to inhibit tumoral growth. Provided are carrier:DNA complexes which comprise cationic polymers or cationic liposomes and DNA encoding at least one anti-angiogenic protein/peptide, optionally together with further DNA encoding a tumor suppressor protein, esp. p53. When administered to a subject bearing a tumor, the complexes can inhibit growth of the tumor.

IT 226938-38-1P, Endostatin (human fragment)

RL: BAC (Biological activity or effector, except adverse); BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(amino acid sequence; delivery of anti-angiogenic genes to a tumor in vivo and their use in gene therapy)

REFERENCE COUNT:

6

REFERENCE(S):

- (1) Chiron Viagene Inc; WO 9621416 A 1996 CAPLUS
- (2) Lesoon-Wood, L; Human Gene Therapy 1995, V6(4), P395 CAPLUS
- (3) Mixson, A; EP 0819758 A 1998 CAPLUS
- (4) The Children's Medical Center Corporation; WO 9529242 A 1995 CAPLUS
- (5) Weinstat-Saslow, D; Cancer Research 1994, V54, P6504 CAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 13 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1999:354395 CAPLUS

DOCUMENT NUMBER:

130:357142

TITLE:

Anti-angiogenic gene therapy vectors and their use in treating angiogenesis-related diseases

INVENTOR (S):

Leboulch, Philippe; Pawliuk, Robert James;

Bachelot, Thomas

PATENT ASSIGNEE(S):

Genetix Pharmaceuticals, Inc., USA; Searcher: Shears 308-4994

Massachusetts Institute of Technology

SOURCE: PCT Int. Appl., 83 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT	NO.	KIN	D DA	Έ		A	PPLI	CATI	ON NO	o. :	DATE		
	- 					-		- -					
WO 992	5480	A1	. 199	90603		W	0 19	98-U	5249	50	1998:	1120	
W:	AL, AM,	AT,	AU, A	, BA,	BB,	BG,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,
	DE, DK,	EE,	ES, F	, GB,	GE,	GH,	GM,	HR,	HU,	ID,	IL,	IS,	JP,
	KE, KG,	KP,	KR, KZ	, LC,	LK,	LR,	LS,	LT,	LU,	LV,	MD,	MG,	MK,
	MN, MW,	MX,	NO, NZ	, PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,
	TJ, TM,	TR,	TT, UA	, UG,	UZ,	VN,	ΥU,	ZW,	AM,	ΑZ,	BY,	KG,	KZ,
	MD, RU,	ТJ,	TM										
RW	GH, GM,	KΕ,	LS, MV	i, SD,	SZ,	ŪĠ,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,
	ES, FI,	FR,	GB, GF	, IE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,
	CG, CI,	CM,	GA, GN	r, GW,	ML,	MR,	NE,	SN,	TD,	TG			
AU 991	985	A1	199	90615		A	U 19	99-1	5985		1998:	1120	
PRIORITY AP	PLN. INFO).:				US 1997-975424 19971120							
						W	0 19	98-U	5249!	50	1998:	1120	

- AB A method for inhibiting tumor growth in a human patient harboring a solid tumor, said method comprising administering to said patient a nucleic acid mol. which expresses in said patient an anti-angiogenic polypeptide selected from the group consisting of human angiostatin, murine angiostatin, human endostatin, murine endostatin, and angiogenesis-inhibiting fragments thereof, wherein expression of the anti-angiogenic polypeptide in the patient inhibits angiogenesis in the vicinity of the tumor and/or systemically by diffusion of the recombinant protein to the vascular compartment from secreting transduced cells, thereby inhibiting its growth.
- IT 224308-23-0, Endostatin mouse

RL: BSU (Biological study, unclassified); BIOL (Biological study) (nucleic acid encoding; anti-angiogenic gene therapy vectors and their use in treating angiogenesis-related diseases)

REFERENCE COUNT:

10

REFERENCE(S):

- (1) Abbott Laboratories; WO 97/41824 A2 1997 CAPLUS
- (2) O'Reilly; US 5792845 A 1998 CAPLUS
- (3) O'Reilly; Cell 1997, V88, P277 CAPLUS
- (4) O'Reilly; Nature Medicine 1996, V2(6), P689 CAPLUS
- (5) Rhone-Poulenc Rorer; WO 98/49321 A2 1998 CAPLUS

ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 14 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1999:134195 CAPLUS

DOCUMENT NUMBER:

130:332418

TITLE:

Antiangiogenic Activity of Restin, NC10 Domain of Human Collagen XV: Comparison to Endostatin

AUTHOR (S):

Ramchandran, Ramani; Dhanabal, Mohanraj; Volk, Ruediger; Waterman, Matthew J. F.; Segal, Mark; Lu, Hua; Knebelmann, Bertrand; Sukhatme, Vikas

Р.

CORPORATE SOURCE:

Renal Div., Dep. Med., Beth Israel Deaconess Med. Cent., Harvard Med. Sch., Boston, MA,

02215, USA

SOURCE:

Biochem. Biophys. Res. Commun. (1999), 255(3),

735-739

CODEN: BBRCA9; ISSN: 0006-291X

PUBLISHER:

Academic Press

DOCUMENT TYPE:

Journal

LANGUAGE:

English

AB Based on a homol. search with endostatin, the C-terminus 185 aa of collagen XVIII, the authors report the cloning, expression, and antiangiogenic activity of a 22 kDa human collagen XV fragment, that the authors have named restin. Restin was expressed in the prokaryotic pET expression system. The authors have shown that restin inhibits the migration of endothelial cells in vitro but has no effect on the proliferation of these cells. A polyclonal antibody raised against endostatin cross-reacted with restin. Systemic administration of restin suppressed the growth of tumors in a xenograft renal carcinoma model. (c) 1999 Academic Press.

IT 224308-23-0

RL: BAC (Biological activity or effector, except adverse); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (amino acid sequence; antiangiogenic activity of NC10 domain of human collagen XV restin in comparison to endostatin in relation to antitumor activity)

REFERENCE COUNT:

13

REFERENCE(S):

- (1) Angiolillo, A; J Exp Med 1995, V182, P155 CAPLUS
- (2) Boehm, T; Nature 1997, V390, P404 CAPLUS
- (3) Corpet, F; Nucleic Acids Res 1988, V16, P10881 CAPLUS
- (5) Folkman, J; Mol Med 1995, V1, P120 CAPLUS
- (6) Folkman, J; Science 1987, V235, P442 CAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 15 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1997:510181 CAPLUS

DOCUMENT NUMBER:

127:146308

TITLE:

Collagen type .alpha.1 (XVIII): a novel member of the collagen family and its properties and

uses

Searcher

Shears 308-4994

INVENTOR(S):

Olsen, Bjorn R.; Oh, Suk P.

PATENT ASSIGNEE(S):

President and Fellows of Harvard College, USA

SOURCE:

U.S., 35 pp.

CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ---------_____ US 5643783 19970701 US 1993-159784 Α 19931201

AB A novel collagen, type .alpha.1 (XVIII), is identified and characterized for therapeutic use. The protein can be conjugated with glycosaminoglycans and used as a carrier for proteins such as fibroblast growth factor (no data) or it can be used as a connective tissue filler in plastic surgery and dermatol. (no data). CDNAs encoding the mouse and human proteins are cloned and antibodies are raised against it. A cDNA for the mouse type .alpha.1 (XVIII) was cloned by screening com. fetal mouse cDNA libraries with probes derived from a type XII collagen. These clones were used to identify a clone for a human type .alpha.1 (XVIII) collagen. domain organization of type .alpha.1 (XVIII) and of type .alpha.1(XV) is different from that of other collagens.

IT 193227-36-0

> RL: BSU (Biological study, unclassified); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (amino acid sequence; collagen type .alpha.1 (XVIII): novel member of collagen family and its properties and uses)

ANSWER 16 OF 19 CAPLUS COPYRIGHT 2001 ACS L2

ACCESSION NUMBER:

1996:185568 CAPLUS

DOCUMENT NUMBER:

124:252313

TITLE:

Characterization of the mouse gene for the .alpha.1 chain of type XVIII collagen (Col18a1)

reveals that the three variant N-terminal polypeptide forms are transcribed from two

widely separated promoters

AUTHOR (S):

Rehn, Marko; Hintikka, Elina; Pihlajaniemi,

Taina

CORPORATE SOURCE:

Collagen Res. Unit, Univ. Oulu, Oulu, FIN-90220,

SOURCE:

Genomics (1996), 32(3), 436-46 CODEN: GNMCEP; ISSN: 0888-7543

DOCUMENT TYPE:

Journal

LANGUAGE:

English

ΔR

The mouse gene for the .alpha.1 chain of type XVIII collagen (Col18a1) is more than 102 kb and consists of 43 exons. Type XVIII collagen transcripts encode polypeptides that differ with respect to

:

Searcher

Shears

308-4994

three variant N-terminal noncollagenous domains that are 301 (NC1-301), 517 (NC1-517), or 764 (NC1-764) residues in length. Characterization of genomic clones revealed that the three variant NC1 domains result from the use of two alternative promoters, sepd. by a distance of 50 kb. The upstream promoter, promoter 1, directs the synthesis of the NC1-301 domain in conjunction with exons 1 and 2, whereas the downstream promoter, promoter 2, directs that of the NC1-517 and NC1-764 domains in conjunction with exon 3, with the latter two variants differing with respect to alternative splicing of the exon 3 sequences. Exons 4-9 encode a portion of the NC1 domain shared by all three polypeptide variants, and exons 9-43 encode the common collagenous and C-terminal noncollagenous sequences. The marked differences previously obsd. in the expression of variant type XVIII collagen transcripts in mouse tissues thus result from tissue-specific use of these two promoters. 175337-10-7 175337-11-8 175337-12-9

RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)

(amino acid sequence; characterization of the mouse gene for the .alpha.1 chain of type XVIII collagen (Coll8a1) reveals that the three variant N-terminal polypeptide forms are transcribed from two widely sepd. promoters)

L2 ANSWER 17 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1994:476556 CAPLUS

DOCUMENT NUMBER: 121:76556

ΙT

TITLE: Isolation and sequencing of cDNAs for proteins

with multiple domains of Gly-Xaa-Yaa repeats identify a distinct family of collagenous

proteins

AUTHOR(S): Oh, Suk Paul; Kamagata, Yusuke; Muragaki,

Yasuteru; Timmons, Sheila; Ooshima, Akira;

Olsen, Bjorn R.

CORPORATE SOURCE: Dep. Cell Biology, Harvard Med. Sch., Boston,

MA, 02115, USA

SOURCE: Proc. Natl. Acad. Sci. U. S. A. (1994), 91(10),

4229-33

CODEN: PNASA6; ISSN: 0027-8424

DOCUMENT TYPE: Journal LANGUAGE: English

AB Overlapping mouse cDNAs were isolated encoding a collagenous polypeptide designated .alpha.1(XVIII) collagen. Nucleotide sequence anal. shows that .alpha.1(XVIII) collagen contains 10 triple-helical domains sepd. and flanked by non-triple-helical regions. Within the non-triple-helical regions, there are several Ser-Gly-contg. sequences that conform to consensus sequences for glycosaminoglycan attachment sites in proteoglycan core proteins. Northern blots show that .alpha.1(XVIII) transcripts are present in multiple organs, with the highest levels in liver, lung, and kidney.

Also, overlapping cDNAs were isolated encoding human .alpha.1(XV) collagen, and their sequence extends a published partial .alpha.1(XV) sequence to the 3' end. Comparison of the .alpha.1(XV) and .alpha.1(XVIII) sequences reveals a striking similarity in the lengths of the 6 most C-terminal triple-helical domains. In addn., within the carboxy non-triple-helical domain NC1 of the 2 chains, a region of 177 amino acid residues shows .apprx.60% identity at the amino acid level. It is suggested, therefore, that .alpha.1(XV) and .alpha.1(XVIII) collagens are structurally related. Their structure is different from that of other known collagen types. Evidently, that they belong to a subfamily of extracellular matrix proteins and the designation multiplexins (for protein with multiple triple-helix domains and interruptions) is suggested for members of this subfamily.

156655-88-8, Collagen (mouse clones mc19/mcE4/mc3 ΙT

.alpha.1(SVIII) - chain fragment)

RL: BIOL (Biological study)

(amino acid sequence and domain structure of)

ANSWER 18 OF 19 CAPLUS COPYRIGHT 2001 ACS L2

ACCESSION NUMBER: 1994:428272 CAPLUS

DOCUMENT NUMBER: 121:28272

Primary structure of the .alpha.1 chain of mouse TITLE:

> type XVIII collagen, partial structure of the corresponding gene, and comparison of the .alpha.1(XVIII) chain with its homolog, the

.alpha.1(XV) collagen chain

Rehn, Marko; Hintikka, Elina; Pihlajaniemi, AUTHOR (S):

Taina

Biocent., Univ. Oulu, Oulu, FIN-90220, Finland CORPORATE SOURCE: SOURCE:

J. Biol. Chem. (1994), 269(19), 13929-35

CODEN: JBCHA3; ISSN: 0021-9258

DOCUMENT TYPE: Journal LANGUAGE: English

The authors have isolated cDNAs that complete the elucidation of the AB primary structure of the mouse .alpha.1(XVIII) collagen chain, a polypeptide homologous to the .alpha.1(XV) collagen chain. 1315-residue .alpha.1(XVIII) chain includes a 25-residue signal peptide, a 301-residue NH2-terminal non-collagenous domain (NC1), a 674-residue collagenous sequence with nine interruptions of 10-24 residues, and a 315-residue COOH-terminal noncollagenous domain (NC11). Seven of the collagenous domains and both flanking noncollagenous domains share homol. with the .alpha.1(XV) chain. The COOH-terminal noncollagenous domains are unique to the .alpha.1(XVIII) and .alpha.1(XV) chains, and they contain a homologous beginning, a variable portion, and a highly homologous COOH-terminal half with 4 conserved cysteines. The differences in the collagenous sequences probably preclude the existence of the two chains in the same mol., however. A 12.5-kilobase pair genomic

Searcher Shears 308-4994 :

sequence was found to contain the 12 extreme 3'-exons of the .alpha.1(XVIII) gene, covering 40% of the coding sequences. Exons start with either a complete codon or a split codon for the glycines of Gly-Xaa-Yaa repeats, and seven exons completely cover the NC11 domain. Comparison of the sequences encoded by these seven exons with the corresponding region of the .alpha.1(XV) gene indicated conserved exon-intron organization, suggesting that the two genes derived from a common ancestor.

IT 155982-66-4, Collagen .alpha.1 chain (mouse clone MM-103

gene COL18A1 type XVIII C-terminal fragment)

RL: PRP (Properties)

(amino acid sequence of)

L2 ANSWER 19 OF 19 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1994:292270 CAPLUS

DOCUMENT NUMBER: 120:292270

TITLE: Identification of a novel collagen chain

represented by extensive interruptions in the

triple-helical region

AUTHOR(S): Abe, Nobuhiro; Muragaki, Yasuteru; Hoshioka,

Hidekatsu; Inoue, Hajime; Ninomiya, Yoshifumi Med. Sch., Okayama Univ., Okayama, 700, Japan

CORPORATE SOURCE: Med. Sch., Okayama Univ., Okayama, 700, Japan SOURCE: Biochem. Biophys. Res. Commun. (1993), 196(2),

576-82

CODEN: BBRCA9; ISSN: 0006-291X

DOCUMENT TYPE: Journal LANGUAGE: English

AB The authors have isolated mouse cDNA clones that encode a newly discovered collagenous polypeptide. Four overlapping cDNA clones contained an open reading frame of 1140 amino acid residues of collagenous and non-collagenous domains. The predicted polypeptide consists of 10 collagenous domains of various length that are interrupted by 11 noncollagenous domains. The overall structural arrangement differed significantly from reported collagen chains of 17 different types. Northern-blot analyses showed hybridization of the cDNA to 5.6kb and 4.8kb mRNA species in liver and kidney that are due to utilization of the two poly (A) signals.

IT 152924-73-7

RL: PRP (Properties)

(amino acid sequence of, extensive interruptions in the triple-helical region in relation to)

E18 THROUGH E34 ASSIGNED

FILE 'REGISTRY' ENTERED AT 10:59:43 ON 06 APR 2001

L3 17 SEA FILE=REGISTRY ABB=ON PLU=ON (224308-23-0/BI OR 226938-38-1/BI OR 259789-72-5/BI OR 326629-20-3/BI OR 152924-73-7/BI OR 155982-66-4/BI OR 156655-88-8/BI OR 175337-10-7/BI OR 175337-11-8/BI OR 175337-12-9/BI OR

193227-36-0/BI OR 255811-03-1/BI OR 303042-57-1/BI OR

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303113-25-9/BI OR 304489-40-5/BI OR 307924-80-7/BI OR
                326948-44-1/BI)
            17 L1 AND L3
L4
=> d 1-17 .bevreg1
     ANSWER 1 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     326948-44-1 REGISTRY
     13: PN: WOO112830 SEQID: 12 unclaimed protein (9CI) (CA INDEX NAME)
CN
CI
    MAN
SQL
    207
        1 METDTLLLWV LLLWVPGSTG DAAHTHQDFQ PVLHLVALNT PLSGGMRGIR
SEQ
        51 GADFQCFQQA RAVGLSGTFR AFLSSRLQDL YSIVRRADRG SVPIVNLKDE
       101 VLSPSWDSLF SGSQGQVQPG ARIFSFDGRD VLRHPAWPQK SVWHGSDPSG
       151 RRLMESYCET WRTETTGATG QASSLLSGRL LEQKAASCHN SYIVLCIENS
       201 FMTSFSK
HITS AT:
          191-198
REFERENCE
          1: 134:188970
    ANSWER 2 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     326629-20-3 REGISTRY
    Protein (mouse strain C57BL/6J clone 3200001M10 160-amino acid)
CN
          (CA INDEX NAME)
     (9CI)
OTHER NAMES:
CN
    GenBank AK014292-derived protein GI 12852042
CI
SQL
    160
        1 GIRGADFQCF QQARAVGLSG TFRAFLSSRL QDLYSIVRRA DRGSVPIVNL
SEQ
        51 KDEVLSPSWD SLFSGSOGOL OPGARIFSFD GRDVLRHPAW PQKSVWHGSD
       101 PSGRRLMESY CETWRTETTG ATGQASSLLS GRLLEQKAAS CHNSYIVLCI
       151 ENSFMTSFSK
HITS AT:
           144-151
    ANSWER 3 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
    307924-80-7 REGISTRY
RN
    L-Methionine, L-.alpha.-glutamyl-L-seryl-L-tyrosyl-L-cysteinyl-L-
     .alpha.-qlutamyl-L-threonyl-L-tryptophyl-L-arginyl-L-threonyl-L-
     .alpha.-glutamyl-L-threonyl-L-threonylglycyl-L-alanyl-L-
     threonylglycyl-L-glutaminyl-L-alanyl-L-seryl-L-seryl-L-leucyl-L-
     leucyl-L-serylglycyl-L-arginyl-L-leucyl-L-leucyl-L-.alpha.-glutamyl-
     L-glutaminyl-L-lysyl-L-alanyl-L-alanyl-L-seryl-L-cysteinyl-L-
                          Searcher
                                             Shears
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histidyl-L-asparaginyl-L-seryl-L-tyrosyl-L-isoleucyl-L-valyl-L-
     leucyl-L-cysteinyl-L-isoleucyl-L-.alpha.-glutamyl-L-asparaginyl-L-
     seryl-L-phenylalanyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
     11: PN: WO0067771 SEQID: 21 claimed protein
CN
CI
     MAN
    48
SQL
         1 ESYCETWRTE TTGATGQASS LLSGRLLEQK AASCHNSYIV LCIENSFM
SEQ
HITS AT:
           37-44
REFERENCE
          1: 134:505
     ANSWER 4 OF 17 REGISTRY COPYRIGHT 2001 ACS
T.4
RN
     304489-40-5 REGISTRY
     Peptide (synthetic clone H6PQE60 histidine tag) fusion protein with
CN
     endostatin (mouse) (9CI) (CA INDEX NAME)
OTHER NAMES:
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CN
CN
     49: PN: WO0064247 SEQID: 13 claimed protein
     MAN
CI
SQL 191
         1 MHHHHHHHH QDFQPVLHLV ALNTPLSGGM RGIRGADFQC FQQARAVGLS
SEQ
        51 GTFRAFLSSR LQDLYSIVRR ADRGSVPIVN LKDEVLSPSW DSLFSGSQGQ
       101 LQPGARIFSF DGRDVLRHPA WPQKSVWHGS DPSGRRLMES YCETWRTETT
       151 GATGQASSLL SGRLLEQKAA SCHNSYIVLC IENSFMTSFS K
                                     _____ __
HITS AT:
           175-182
REFERENCE
           1: 133:349140
    ANSWER 5 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     303113-25-9 REGISTRY
     L-Lysine, L-cysteinyl-L-.alpha.-glutamyl-L-threonyl-L-tryptophyl-L-
CN
     arginyl-L-threonyl-L-.alpha.-glutamyl-L-threonyl-L-threonylglycyl-L-
     alanyl-L-threonylglycyl-L-glutaminyl-L-alanyl-L-seryl-L-seryl-L-
     leucyl-L-leucyl-L-serylglycyl-L-arginyl-L-leucyl-L-leucyl-L-.alpha.-
     glutamyl-L-glutaminyl-L-lysyl-L-alanyl-L-alanyl-L-seryl-L-cysteinyl-
     L-histidyl-L-asparaginyl-L-seryl-L-tyrosyl-L-isoleucyl-L-valyl-L-
     leucyl-S-(1,1-dimethylethyl)-L-cysteinyl-L-isoleucyl-L-.alpha.-
     glutamyl-L-asparaginyl-L-seryl-L-phenylalanyl-L-methionyl-L-threonyl-
     L-seryl-L-phenylalanyl-L-seryl-, cyclic (1.fwdarw.31)-disulfide
     (9CI)
           (CA INDEX NAME)
OTHER NAMES:
     4: PN: WO0063249 SEQID: 4 claimed protein
CN
CI
     MAN
SQL 50
                          Searcher
                                             Shears
                                                       308-4994
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1 CETWRTETTG ATGQASSLLS GRLLEQKAAS CHNSYIVLCI ENSFMTSFSK
SEQ
HITS AT:
           34-41
REFERENCE
            1: 133:329588
     ANSWER 6 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
     303042-57-1 REGISTRY
RN
     L-Lysine, S-(triphenylmethyl)-L-cysteinyl-L-.alpha.-glutamyl-O-(1,1-
CN
     dimethylethyl) -L-threonyl-1-[(1,1-dimethylethoxy)carbonyl]-L-
     tryptophyl-N5-[[[(2,3-dihydro-2,2,4,6,7-pentamethyl-5-
     benzofuranyl) sulfonyl] amino] iminomethyl] -L-ornithyl-O-(1,1-
     dimethylethyl)-L-threonyl-L-.alpha.-glutamyl-O-(1,1-dimethylethyl)-L-
     threonyl-O-(1,1-dimethylethyl)-L-threonylglycyl-L-alanyl-O-(1,1-
     dimethylethyl) -L-threonylglycyl-L-glutaminyl-L-alanyl-O-(1,1-
     dimethylethyl) -L-seryl-O-(1,1-dimethylethyl) -L-seryl-L-leucyl-L-
     leucyl-O-(1,1-dimethylethyl)-L-serylglycyl-N5-[[[(2,3-dihydro-
     2,2,4,6,7-pentamethyl-5-benzofuranyl)sulfonyl]amino]iminomethyl]-L-
     ornithyl-L-leucyl-L-leucyl-L-.alpha.-glutamyl-L-glutaminyl-N6-[(1,1-
     dimethylethoxy) carbonyl]-L-lysyl-L-alanyl-L-alanyl-O-(1,1-
     dimethylethyl) -L-seryl-S-(triphenylmethyl) -L-cysteinyl-1-
     (triphenylmethyl) -L-histidyl-L-asparaginyl-O-(1,1-dimethylethyl) -L-
     seryl-O-(1,1-dimethylethyl)-L-tyrosyl-L-isoleucyl-L-valyl-L-leucyl-S-
     (1,1-dimethylethyl)-L-cysteinyl-L-isoleucyl-L-.alpha.-glutamyl-L-
     asparaginyl-O-(1,1-dimethylethyl)-L-seryl-L-phenylalanyl-L-methionyl-
     O-(1,1-dimethylethyl)-L-threonyl-O-(1,1-dimethylethyl)-L-seryl-L-
     phenylalanyl-O-(1,1-dimethylethyl)-L-seryl-, 2,7,25,41-tetrakis(1,1-
     dimethylethyl) ester (9CI) (CA INDEX NAME)
OTHER NAMES:
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CN
CI
     MAN
SOL
     50
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HITS AT:
           34-41
            1: 133:329588
REFERENCE
     ANSWER 7 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     259789-72-5 REGISTRY
                               (CA INDEX NAME)
     Endostatin (mouse) (9CI)
CN
OTHER NAMES:
CN
     18: PN: WO0011033 SEQID: 18 claimed protein
CI
     MAN
SQL
    184
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SEQ
                                             Shears
                                                        308-4994
                          Searcher
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51 SSRLQDLYSI VRRADRGSVP IVNLKDEVLS PSWDSLFSGS QGQVQPGARI
       101 FSFDGRDVLR HPAWPQKSVW HGSDPSGRRL MESYCETWRT ETTGATGQAS
       151 SLLSGRLLEQ KAASCHNSYI VLCIENSFMT SFSK
         168-175
HITS AT:
REFERENCE 1: 133:292674
REFERENCE 2: 132:190503
    ANSWER 8 OF 17 REGISTRY COPYRIGHT 2001 ACS
    255811-03-1 REGISTRY
    4: PN: CN1177005 PAGE: 4 unclaimed sequence (9CI) (CA INDEX NAME)
    MAN
SQL 181
        1 HTHQDFQPVL HLVALNTPLS GGMRGIRGAD FQCFQQARAV GLSGTFRAFL
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       101 FSFDGRDVLR HPAWPQKSVW HGSDPSGRRL MESYCETWRT ETTGATGQAS
       151 SLLSGRLLEQ KAASCHNSYI VLCIENSFMT X
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HITS AT: 168-175
REFERENCE 1: 132:103744
    ANSWER 9 OF 17 REGISTRY COPYRIGHT 2001 ACS
    226938-38-1 REGISTRY
    Endostatin (human fragment) (9CI) (CA INDEX NAME)
OTHER NAMES:
   31: PN: US6080728 SEQID: 36 claimed protein
    Endostatin (synthetic 185-amino acid fragment)
    MAN
SQL 185
        1 MHTHQDFQPV LHLVALNTPL SGGMRGIRGA DFQCFNNARV GLSGTFRAFL
       51 SSRLQDLYSI VRRADRGSVP IVQNLRDEVL SPSWDSLFSG SQGQLQPGAR
      101 IFSFDGRDVL RHPAWPQRSV WHGSDPSGRR LMESYCETWR TETTGATGQA
      151 SSLLSGRLLE QRAASCHDSY IVLCIENSFM TSFSR
HITS AT: 169-176
REFERENCE 1: 133:79332
REFERENCE 2: 131:28626_____
    ANSWER 10 OF 17 REGISTRY COPYRIGHT 2001 ACS
    224308-23-0 REGISTRY
    957-1140-Collagen (mouse clone NA1/NA12/NA2611/NA286 reduced) (9CI)
     (CA INDEX NAME)
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Searcher: Shears 308-4994

L4RN

CN CI

SEO

L4

RN

CN

CN

CN CI/

SEO

T.4

RN

CN

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OTHER NAMES:
ÇN
     4: PN: WO0067771 SEQID: 4 unclaimed protein
CN
     Endostatin (mouse)
CI
     MAN
SQL
    184
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SEO
        51 SSRLODLYSI VRRADRGSVP IVNLKDEVLS PSWDSLFSGS QGQLQPGARI
       101 FSFDGRDVLR HPAWPQKSVW HGSDPSGRRL MESYCETWRT ETTGATGQAS
       151 SLLSGRLLEQ KAASCHNSYI VLCIENSFMT SFSK
                             === ====
HITS AT:
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REFERENCE
          1: 134:505
REFERENCE 2: 131:41277
REFERENCE
          3: 130:357142
REFERENCE
            4: 130:332418
     ANSWER 11 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
     193227-36-0 REGISTRY
RN
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     (CA INDEX NAME)
CI
    MAN
    1288
SQL
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SEQ
       51 KLFFRDFSLL FHVRPATEAA GVLFAITDAA QVVVSLGVKL SEVRDGQQNI
       101 SLLYTEPGAS QTQTGASFRL PAFVGQWTHF ALSVDGGSVA LYVDCEEFQR
      151 VPFARASQGL ELERGAGLFV GQAGTADPDK FQGMISELKV RKTPRVSPVH
      201 CLDEEDDDED RASGDFGSGF EESSKSHKED TSLLPGLPQP PPVTSPPLAG
      251 GSTTEDPRTE ETEEDAAVDS IGAETLPGTG SSGAWDEAIQ NPGRGLIKGG
      301 MKGOKGEPGA OGPPGPAGPO GPAGPVVQSP NSQPVPGAQG PPGPQGPPGK
      351 DGTPGRDGEP GDPGEDGRPG DTGPQGFPGT PGDVGPKGEK GDPGIGPRGP
      401 PGPPGPPGPS FRQDKLTFID MEGSGFSGDI ESLRGPRGFP GPPGPPGVPG
      451 LPGEPGRFGI NGSYAPGPAG LPGVPGKEGP PGFPGPPGPP GPPGKEGPPG
      501 VAGOKGSVGD VGIPGPKGSK GDLGPIGMPG KSGLAGSPGP VGPPGPPGPP
      551 GPPGPGFAAG FDDMEGSGIP LWTTARSSDG LQGPPGSPGL KGDPGVAGLP
      601 GAKGEVGADG AQGIPGPPGR EGAAGSPGPK GEKGMPGEKG NPGKDGVGRP
      651 GLPGPPGPPG PVIYVSSEDK AIVSTPGPEG KPGYAGFPGP AGPKGDLGSK
      701 GEQGLPGPKG EKGEPGTIFS PDGRALGHPQ KGAKGEPGFR GPPGPYGRPG
      751 HKGEIGFPGR PGRPGTNGLK GEKGEPGDAS LGFSMRGLPG PPGPPGPPGP
      801 PGMPIYDSNA FVESGRPGLP GQQGVQGPSG PKGDKGEVGP PGPPGQFPID
      851 LFHLEAEMKG DKGDRGDAGQ KGERGEPGAP GGGFFSSSVP GPPGPPGYPG
      901 IPGPKGESIR GPPGPPGPQG PPGIGYEGRQ GPPGPPGPPG PPSFPGPHRQ
       951 TVSVPGPPGP PGPPGPPGAM GASAGOVRIW ATYQTMLDKI REVPEGWLIF
      1001 VAEREELYVR VRNGFRKVLL EARTALLRGT GNEVAAFQPP LVQLHEGSPY
```

Searcher

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Shears

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1051 TRREYSYSTA RPWRADDILA NPPRLPDRQP YPGVPHHHSS YVHLPPARPT
                                                               INOP Same EN EMI
FOR EVg. 2
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      1151 RAFLSSRLQD LYSIVRRADR GSVPIVNLKD EVLSPSWDSL FSGSQGQVQP
      1201 GARIFSFDGR DVLRHPAWPQ KSVWHGSDPS GRRLMESYCE TWRTETTGAT
      1251 GQASSLLSGR LLEQKAASCH NSYIVLCIEN SFMTSFSK
HITS AT:
           1272-1279
REFERENCE
            1: 127:146308
     ANSWER 12 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     175337-12-9 REGISTRY
    Collagen (mouse clone P13124/GEN19/K17.6 type XVIII gene Col18a1
CN
     .alpha.1-chain 1527-amino acid isoform precursor reduced) (9CI)
     INDEX NAME)
CI
    MAN
SQL
    1527
SEQ
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        51 GPVSKPQNSS PVQSTENPTT HVVPQDGLTE QQTTPASSEL PPEEEEEEDQ
       101 KAGQGGSPAT PAVPIPLVAP AASPDMKEEN VAGVGAKILN VAQGIRSFVQ
       151 LWDEDSTIGH SAGTEVPDSS IPTVLPSPAE LSSAPQGSKT TLWLSSAIPS
       201 SPDAOTTEAG TLAVPTOLPP FOSNLOAPLG RPSAPPDFPE NVAEEVGLLO
       251 LLGDPLPEKI SQIDDPHVGP AYIFGPDSNS GQVAQYHFPK LFFRDFSLLF
       301 HVRPATEAAG VLFAITDAAQ VVVSLGVKLS EVRDGQQNIS LLYTEPGASQ
       351 TQTGASFRLP AFVGQWTHFA LSVDGGSVAL YVDCEEFQRV PFARASQGLE
       401 LERGAGLFVG QAGTADPDKF QGMISELKVR KTPRVSPVHC LDEEDDDEDR
       451 ASGDFGSGFE ESSKSHKEDT SLLPGLPQPP PVTSPPLAGG STTEDPRTEE
       501 TEEDAAVDSI GAETLPGTGS SGAWDEAIQN PGRGLIKGGM KGQKGEPGAQ
      551 GPPGPAGPOG PAGPVVOSPN SOPVPGAOGP PGPOGPPGKD GTPGRDGEPG
       601 DPGEDGRPGD TGPQGFPGTP GDVGPKGEKG DPGIGPRGPP GPPGPPGPSF
       651 RODKLTFIDM EGSGFSGDIE SLRGPRGFPG PPGPPGVPGL PGEPGRFGIN
      701 GSYAPGPAGL PGVPGKEGPP GFPGPPGPPG PPGKEGPPGV AGQKGSVGDV
      751 GIPGPKGSKG DLGPIGMPGK SGLAGSPGPV GPPGPPGPPG PPGPGFAAGF
      801 DDMEGSGIPL WTTARSSDGL QGPPGSPGLK GDPGVAGLPG AKGEVGADGA
      851 QGIPGPPGRE GAAGSPGPKG EKGMPGEKGN PGKDGVGRPG LPGPPGPPGP
      901 VIYVSSEDKA IVSTPGPEGK PGYAGFPGPA GPKGDLGSKG EQGLPGPKGE
      951 KGEPGTIFSP DGRALGHPQK GAKGEPGFRG PPGPYGRPGH KGEIGFPGRP
     1001 GRPGTNGLKG EKGEPGDASL GFSMRGLPGP PGPPGPPGPP GMPIYDSNAF
     1051 VESGRPGLPG QQGVQGPSGP KGDKGEVGPP GPPGQFPIDL FHLEAEMKGD
     1101 KGDRGDAGQK GERGEPGAPG GGFFSSSVPG PPGPPGYPGI PGPKGESIRG
     1151 PPGPPGRQGP PGIGYEGRQG PPGPPGPPGP PSFPGPHRQT VSVPGPPGPP
     1201 GPPGPPGAMG ASAGQVRIWA TYQTMLDKIR EVPEGWLIFV AEREELYVRV
     1251 RNGFRKVLLE ARTALPRGTG NEVAALQPPL VQLHEGSPYT RREYSYSTAR
     1301 PWRADDILAN PPRLPDROPY PGVPHHHSSY VHLPPARPTL SLAHTHODFO
     1351 PVLHLVALNT PLSGGMRGIR GADFQCFQQA RAVGLSGTFR AFLSSRLQDL
     1401 YSIVRRADRG SVPIVNLKDE VLSPSWDSLF SGSQGQLQPG ARIFSFDGRD
     1451 VLRHPAWPQK SVWHGSDPSG RRLMESYCET WRTETTGATG QASSLLSGRL
     1501 LEQKAASCHN SYIVLCIENS FMTSFSK
                                                       308-4994
                         Searcher
                                             Shears
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HITS AT: 1511-1518 REFERENCE 1: 124:252313 L4ANSWER 13 OF 17 REGISTRY COPYRIGHT 2001 ACS 175337-11-8 REGISTRY RN Collagen (mouse clone P13124/GEN19/K17.6 type XVIII gene Col18a1 CN .alpha.1-chain 1774-amino acid isoform precursor reduced) (9CI) INDEX NAME) CI MAN 1774 SQL 1 MAPDPSRRLC LLLLLLSCR LVPASADGNS LSPLNPLVWL WPPKTSDSLE SEQ 51 GPVSKPONSS PVOSTENPTT HVVPODGLTE OOTTPASSEL PPEEEEEEDO 101 KAGQGGSPAT PAVPIPLVAP AASPDMKEEN VAGVGAKILN VAQGIRSFVQ 151 LWDEDSTIGH SAGTEVPDSS IPTVLPSPAE LSSAPQGSKT TLWLSSAIPS 201 SPDAOTTEAG TLAVPTOLPP FOSNLOAPLG RPSAPPDFPG RAFLSSSTDO 251 GSSWGNQEPP RQPQHLEGKG FLPMTARSSQ QHRHSDVHSD IHGHVPLLPL 301 VTGPLVTASL SVHGLLSVPS SDPSGQLSQV AALPGFPGTW VSHVAPSSGT 351 GLSNDSALAG NGSLTSTSRC LPLPPTLTLC SRLGIGHFWL PNHLHHTDSV 401 EVEATVQAWG RFLHTNCHPF LAWFFCLLLA PSCGPGPPPP LPPCRQFCEA 451 LEDECWNYLA GDRLPVVCAS LPSQEDGYCV FIGPAAENVA EEVGLLQLLG 501 DPLPEKISOI DDPHVGPAYI FGPDSNSGOV AOYHFPKLFF RDFSLLFHVR 551 PATEAAGVLF AITDAAQVVV SLGVKLSEVR DGQQNISLLY TEPGASQTQT 601 GASFRLPAFV GQWTHFALSV DGGSVALYVD CEEFQRVPFA RASQGLELER 651 GAGLFVGQAG TADPDKFQGM ISELKVRKTP RVSPVHCLDE EDDDEDRASG 701 DFGSGFEESS KSHKEDTSLL PGLPOPPPVT SPPLAGGSTT EDPRTEETEE 751 DAAVDSIGAE TLPGTGSSGA WDEAIQNPGR GLIKGGMKGQ KGEPGAQGPP 801 GPAGPQGPAG PVVQSPNSQP VPGAQGPPGP QGPPGKDGTP GRDGEPGDPG 851 EDGRPGDTGP QGFPGTPGDV GPKGEKGDPG IGPRGPPGPP GPPGPSFRQD 901 KLTFIDMEGS GFSGDIESLR GPRGFPGPPG PPGVPGLPGE PGRFGINGSY 951 APGPAGLPGV PGKEGPPGFP GPPGPPGPPG KEGPPGVAGQ KGSVGDVGIP 1001 GPKGSKGDLG PIGMPGKSGL AGSPGPVGPP GPPGPPGPPG PGFAAGFDDM 1051 EGSGIPLWTT ARSSDGLOGP PGSPGLKGDP GVAGLPGAKG EVGADGAQGI 1101 PGPPGREGAA GSPGPKGEKG MPGEKGNPGK DGVGRPGLPG PPGPPGPVIY 1151 VSSEDKAIVS TPGPEGKPGY AGFPGPAGPK GDLGSKGEQG LPGPKGEKGE 1201 PGTIFSPDGR ALGHPQKGAK GEPGFRGPPG PYGRPGHKGE IGFPGRPGRP 1251 GTNGLKGEKG EPGDASLGFS MRGLPGPPGP PGPPGPPGMP IYDSNAFVES 1301 GRPGLPGQQG VQGPSGPKGD KGEVGPPGPP GQFPIDLFHL EAEMKGDKGD 1351 RGDAGOKGER GEPGAPGGGF FSSSVPGPPG PPGYPGIPGP KGESIRGPPG 1401 PPGRQGPPGI GYEGRQGPPG PPGPPGPPSF PGPHRQTVSV PGPPGPPGPP 1451 GPPGAMGASA GQVRIWATYQ TMLDKIREVP EGWLIFVAER EELYVRVRNG 1501 FRKVLLEART ALPRGTGNEV AALQPPLVQL HEGSPYTRRE YSYSTARPWR 1551 ADDILANPPR-LPDROPYPGV PHHHSSYVHL PPARPTLSLA HTHODFOPVL 1601 HLVALNTPLS GGMRGIRGAD FOCFOQARAV GLSGTFRAFL SSRLODLYSI 1651/VRRADRGSVP IVNLKDEVLS PSWDSLFSGS QGQLQPGARI FSFDGRDVLR

1701 HPAWPOKSVW HGSDPSGRRL MESYCETWRT ETTGATGOAS SLLSGRLLEO

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308-4994

-Searcher

1751\KAASCHNSYI VLCIENSFMT SFSK

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Searcher

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C-terminal fragment reduced) (9CI) (CA INDEX NAME)
OTHER NAMES:
     Collagen (mouse clones mc19/mcE4/mc3 .alpha.1(SVIII)-chain fragment)
CI
     MAN
SOL
     1288
SEQ
         1 ENVAEEVGLL OLLGDPLPEK ISOIDDPHVG PAYIFGPDSN SGOVAOYHFP
        51 KLFFRDFSLL FHVRPATEAA GVLFAITDAA QVVVSLGVKL SEVRDGOONI
       101 SLLYTEPGAS QTQTGASFRL PAFVGQWTHF ALSVDGGSVA LYVDCEEFQR
       151 VPFARASOGL ELERGAGLFV GOAGTADPDK FOGMISELKV RKTPRVSPVH
       201 CLDEEDDDED RASGDFGSGF EESSKSHKED TSLLPGLPQP PPVTSPPLAG
       251 GSTTEDPRTE ETEEDAAVDS IGAETLPGTG SSGAWDEAIQ NPGRGLIKGG
       301 MKGQKGEPGA QGPPGPAGPQ GPAGPVVQSP NSQPVPGAQG PPGPQGPPGK
       351 DGTPGRDGEP GDPGEDGRPG DTGPQGFPGT PGDVGPKGEK GDPGIGPRGP
       401 PGPPGPPGPS FRQDKLTFID MEGSGFSGDI ESLRGPRGFP GPPGPPGVPG
       451 LPGEPGRFGI NGSYAPGPAG LPGVPGKEGP PGFPGPPGPP GPPGKEGPPG
       501 VAGQKGSVGD VGIPGPKGSK GDLGPIGMPG KSGLAGSPGP VGPPGPPGPP
       551 GPPGPGFAAG FDDMEGSGIP LWTTARSSDG LOGPPGSPGL KGDPGVAGLP
       601 GAKGEVGADG AQGIPGPPGR EGAAGSPGPK GEKGMPGEKG NPGKDGVGRP
       651 GLPGPPGPPG LVIYVSSEDK AIVSTPGPEG KPGYAGFPGP AGPKGDLGSK
       701 GEQGLPGFKG EKGEPGTIFS PDGRRLGHPQ KGAKGEPGFR GPPGPYGRPG
       751 HKGEIGFPGR PGRPGTNGLK GEKGEPGDAS LGFSMRGLPG PPGPPGPPGP
       801 PGMPIYDSNA FVESGRPGLP GQQGVQGPSG PKGDKGEVGP PGPPGQFPID
       851 LFHLEAEMKG DKGDRGDAGQ KGERGEPGAP GGGFFSSSVP GPPGPPGYPG
       901 IPGPKGESIR GPPGPPGPQG PPGIGYEGRQ GPPGPPGPPG PPSFPGPHRQ
       951 TVSVPGPPGP PGPPGPPGAM GASAGOVRIW ATYOTMLDKI REVPEGWLIF
      1001 VAEREELYVR VRNGFRKVLL EARTALLRGT GNEVAAFQPP LVQLHEGSPY
      1051 TRREYSYSTA RPWRADDILA_NPPRLPDRQP YPGVPHHHSS_YVHLPPARPT
      1101 LSLAHTHODF QPVLHLVALN TPLSGGMRGI RGADFQCFQQ ARAVGLSGTF
      1151 AAFLSSRLOD LYSIVRRADR GSVPIVNLKD EVLSPSWDSL FSGSOGOVOP
      1201 GARIFSFDGR DVLRHPAWPQ KSVWHGSDPS GRRLMESYCE TWRTETTGAT
      1251 GQASSLLSGR LLEQKAASCH NSYIVLCIEN SFMTSFSK
          1272-1279
HITS AT:
REFERENCE
           1: 121:76556
L4
    ANSWER 16 OF 17 REGISTRY COPYRIGHT 2001 ACS
RN
    155982-66-4 REGISTRY
    Collagen (mouse clone MM-103 gene COL18A1 type XVIII .alpha.1-chain
    C-terminal fragment reduced) (9CI) (CA INDEX NAME)
OTHER NAMES:
    Collagen .alpha.1 chain (mouse clone MM-103 gene COL18A1 type XVIII
CN
    C-terminal fragment)
CI
    MAN
SQL
    482
SEQ
         1 DSNAFVESGR PGLPGQQGVQ GPSGPKGDKG EVGPPGPPGQ FPIDLFHLEA
        51 EMKGDKGDRG DAGQKGERGE PGAPGGGFFS SSVPGPPGPP GYPGIPGPKG
                          Searcher
                                             Shears
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101 ESIRGPPGPP GROGPPGIGY EGROGPPGPP GPPGPPSFPG PHRQTVSVPG
       151 PPGPPGPPGP PGAMGASAGQ VRIWATYQTM LDKIREVPEG WLIFVAEREE
       201 LYVRVRNGFR KVLLEARTAL PRGTGNEVAA LQPPLVQLHE GSPYTRREYS
       251 YSTARPWRAD DILANPPRLP DROPYPGVPH HHSSYVHLPP ARPTLSLAHT
       301 HQDFQPVLHL VALNTPLSGG MRGIRGADFQ CFQQARAVGL SGTFRAFLSS
       351/RLQDLYSIVR RADRGSVPIV NLKDEVLSPS WDSLFSGSQG QLQPGARIFS
       401/FDGRDVLRHP AWPOKSVWHG SDPSGRRLME SYCETWRTET TGATGQASSL
       451 LSGRLLEOKA ASCHNSYIVL CIENSFMTSF SK
                          _========
HITS AT:
           466-473
REFERENCE 1: 121:28272
     ANSWER 17 OF 17 REGISTRY COPYRIGHT 2001 ACS
L4
RN
     152924-73-7 REGISTRY
     Collagen (mouse clone NA1/NA12/NA2611/NA286 reduced) (9CI) (CA
CN
CI
     MAN
SQL
    1140
SEQ
         1 QRVPFARASQ GLELERGAGL FVGQAGTADP DKFQGMISEL KVRKTPRVSP
        51 VHCLDEEDDD EDRASGDFGS GFEESSKSHK EDTSLLPGLP QPPPVTSPPL
       101 AGGSTTEDPR TEETEEDAAV DSIGAETLPG TGSSGAWDEA IQNPGRGLIK
       151 GGMKGOKGEP GAOGPPGPAG POGPAGPVVO SPNSQPVPGA QGPPGPQGPP
       201 GKDGTPGRDG EPGDPGEDGR PGDTGPQGFP GTPGDVGPKG EKGDPGIGPR
       251 GPPGPPGPPG PSFRQDKLTF IDMEGSGFSG DIESLRGPRG FPGPPGPPGV
       301 PGLPGEPGRF GINGSYAPGP AGLPGVPGKE GPPGFPGPPG PPGPPGKEGP
       351 PGVAGOKGSV GDVGIPGPKG SKGDLGPIGM PGKSGLAGSP GPVGPPGPPG
       401 PPGPPGPGFA AGFDDMEGSG IPLWTTARSS DGLQGPPGSP GLKGDPGVAG
       451 LPGAKGEVGA DGAQGIPGPP GREGAAGSPG PKGEKGMPGE KGNPGKDGVG
       501 RPGLPGPPGP PGPVIYVSSE DKAIVSTPGP EGKPGYAGFP GPAGPKGDLG
       551 SKGEQGLPGP KGEKGEPGTI FSPDGRALGH PQKGAKGEPG FRGPPGPYGR
       601 PGHKGEIGFP GRPGRPGTNG LKGEKGEPGD ASLGFSMRGL PGPPGPPGPP
       651 GPPGMPIYDS NAFVESGRPG LPGQQGVQGP SGPKGDKGEV GPPGPPGQFP
       701 IDLFHLEAEM KGDKGDRGDA GQKGERGEPG APGGGFFSSS VPGPPGPPGY
       751 PGIPGPKGES IRGPPGPPGP QGPPGIGYEG RQGPPGPPGP PGPPSFPGPH
       801 ROTVSVPGPP GPPGPPGPPG AMGASAGQVR IWATYQTMLD KIREVPEGWL
       851 IFVAEREELY VRVRNGFRKV LLEARTALPR GTGNEVAALQ PPLVQLHEGS
       901 PYTRREYSYS TARPWRADDI LANPPRLPDR QPYPGVPHHH_SSYVHLPPAR
       951 PTLSLAHTHO DFOPVLHLVA LNTPLSGGMR GIRGADFOCF QQARAVGLSG
      1001/TFRAFLSSRL QDLYSIVRRA DRGSVPIVNL KDEVLSPSWD SLFSGSQGQL
      1051 OPGARIFSFD GRDVLRHPAW POKSVWHGSD PSGRRLMESY CETWRTETTG
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REFERENCE
     FILE 'CAPLUS' ENTERED AT 11:00:29 ON 06 APR 2001
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Searcher :

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308-4994

L5 1344 S EM1 OR EMI OR EM(W) (I OR 1) - Key terms
L6 2 S L5 AND (ENDOSTATIN OR ENDO STATIN) - Claims 2-4 \$ 11

L7 1 S L6 NOT L2

L7 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1999:388288 CAPLUS

DOCUMENT NUMBER: 131:39759

TITLE: Restin and apomigren fragments of human collagen

type XV .alpha.1 chain and their anti-angiogenic

activities

INVENTOR(S): Sukhatme, Vikas P.

PATENT ASSIGNEE(S): Beth Israel Deaconess Medical Center, USA

SOURCE: PCT Int. Appl., 94 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE WO 1998-US26058 19981208 WO 9929856 A1 19990617 W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, US, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG AU 9918088 A1 19990628 AU 1999-18088 19981208 EP 1998-962966 20000927 EP 1037985 **A**1 19981208 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI PRIORITY APPLN. INFO.: US 1997-67888 19971208 US 1998-82663 19980422 US 1998-108536 19981116 WO 1998-US26058 19981208

AB The invention relates to restin, a novel anti-angiogenic protein is described, as well as its fragment, designated apomigren. Restin is a proteolytic fragment of the C-terminal fragment of the NC10 domain of the .alpha.l chain of human collagen type XV. Apomigren is a fragment of restin, and comprises the C-terminal 85 residues of restin,. Methods for expression of the proteins at high titer are also described. Restin inhibits the migration of endothelial cells in vitro and suppresses the growth of tumors in a xenograft renal carcinoma model. Apomigren has anti-angiogenic activity equal or superior to that of endostatin.

REFERENCE COUNT:

REFERENCE(S):

- (1) Bachelot; Proceedings of the 89th Annual Meeting of the American Association for Cancer Research 1998, V39, P271
- (2) Childrens Medical Center; WO 9715666 A 1997 **CAPLUS**
- (3) Ramchandran, R; Biochem Biophys Res Comm 1999, V255, P735 CAPLUS
- (4) Rehn, M; J Biol Chem 1994, V269(19), P13929 CAPLUS
- (6) Searle, G; WO 9916899 A 1999 CAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT

(FILE 'MEDLINE, BIOSIS, EMBASE, WPIDS, CONFSCI, SCISEARCH, JICST-EPLUS, JAPIO' ENTERED AT 11:01:47 ON 06 APR 2001)

L8

1 S L6

L8 ANSWER 1 OF 1 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD

ACCESSION NUMBER: 1999-385604 [32] WPIDS

1999-394974 [33]; 1999-404943 [34]

TITLE:

Mutant endostatin having anti-angiogenic

activity.

DERWENT CLASS:

B04 D16

INVENTOR(S):

SUKHATME, V P

PATENT ASSIGNEE(S): (BETH-N) BETH ISRAEL DEACONESS MEDICAL CENT

COUNTRY COUNT:

85

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA -----

A1 19990617 (199932)* EN 105

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW

W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG US UZ VN YU ZW

AU 9917180 A 19990628 (199946)

EP 1037983 A1 20000927 (200048) EN

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

APPLICATION DETAILS:

PATENT NO	KIND		APPLICATION					
WO 9929855	A1	·	wo	1998-US26057	19981208			
AU 9917180	Α		ΑU	1999-17180	19981208			
EP 1037983	A1		EP	1998-962006	19981208			
		Searcher	:	Shears	308-4994			

WO 1998-US26057 19981208

FILING DETAILS:

PRIORITY APPLN. INFO: US 1998-108536 19981116; US 1997-67888 19971208; US 1998-82663 19980422

AN 1999-385604 [32] WPIDS

CR 1999-394974 [33]; 1999-404943 [34]

AB WO 9929855 A UPAB: 20001001

NOVELTY - A mutant endostatin (EM) having anti-angiogenic activity comprising a C-terminal sequence (I), is new.

DETAILED DESCRIPTION - An isolated anti-angiogenic peptide, where the C-terminal comprises the amino acid sequence SYIVLCIE (I). INDEPENDENT CLAIMS are also included for the following:

(a) an isolated polynucleotide amplified by the following primers (P1), and (P2):

TTCCATATGCATACTCATCAGGACTTTCAGGCA (P1); and TTAGCGGCCGCCTACTCAATGCAGAGGACGATGTA (P2);

- (b) a host cell transformed with a polynucleotide, encodingEM1, operably linked to an expression control sequence;
 - (c) production of EM1;
- (d) a fusion protein comprising two or more proteins and also comprising EM1;
 - (e) a process for providing a mammal with EM1;
- (f) producing an isolated polynucleotide which hybridizes under moderate stringency;
 - (g) an EM1 polynucleotide isolated by (f);
 - (h) antibodies to EM1; and
 - (i) a mutant, derivative, analogue or homologue of EM1

ACTIVITY - Anti-angiogenic; cytostatic.

MECHANISM OF ACTION - None given.

USE - Compositions comprising EM1 or fusion proteins comprising EM1, are useful for treating diseases characterized by angiogenic activity, such as angiogenesis-dependent cancers, benign tumors, rheumatoid arthritis, psoriasis, ocular angiogenesis, Osler-Webber Syndrome, myocardial angiogenesis, plaque neovascularization, telangiectasia, hemophiliac joints, angiofibroma, wound granulation, intestinal adhesions, atherosclerosis, scleroderma, hypertrophic scars, cat scratch disease, Helicobacter pylori ulcers, dialysis graft vascular access stenosis, contraception and obesity. In particular, the diseases treatable by EM1 comprise cancer, especially renal cancer. The methods provide a means for introducing EM1 into

mammalian cells via gene therapy, for production of EM1 via recombinant means, as well as recombinant production of the EM1 protein. (All claimed).

ADVANTAGE - EM1 performs as well or better than whole endostatin. In a nude mouse model, growth of renal cell cancer (RCC) was suppressed by systemic administration of EM1 at a rate of 20 mg/kg body weight. Use of EM1 is advantageous for treatment of angiogenic diseases in that increasingly smaller peptides are more potent on a weight basis, and may be able to better penetrate tissues.

Dwg.20/26

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FILE 'HOME' ENTERED AT 11:04:41 ON 06 APR 2001